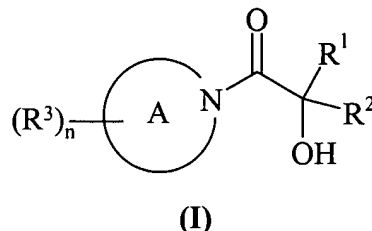


**IN THE CLAIMS:**

Claim 1 (currently amended): A compound of formula (I):



wherein:

**Ring A** is piperazinyl substituted on nitrogen by  $R^4$ -D-;

$R^1$  and  $R^2$  are independently  $C_k$ alkyl optionally substituted by 1 to  $2k+1$  atoms selected from fluoro and chloro wherein  $k$  is 1-3;

or  $R^1$  and  $R^2$  together with the carbon atom to which they are attached, form a  $C_m$ cycloalkyl ring optionally substituted by 1 to  $2m-2$  fluorine atoms wherein  $m$  is 3-5;

$R^3$  is a substituent on carbon and is halo, hydroxy, cyano, formyl, amino, nitro, carboxy, carbamoyl, ureido, thiol, sulphamoyl or  $R^5$ -E-;

$R^4$  is  $C_{1-6}$ alkyl, phenyl or a heterocyclic group, wherein in  $R^4$  any  $C_{1-6}$ alkyl, phenyl or heterocyclic group groups (on a ring carbon) may be optionally substituted by one or more  $R^6$  and if said heterocyclic group contains an NH moiety that nitrogen may be optionally substituted by a group selected from  $R^8$ ;

D is  $-N(R^9)C(O)-$ ,  $-S(O)_2-$  or  $-NS(O)_2-$ ;

$R^5$  is  $C_{1-6}$ alkyl,  $C_{2-6}$ alkenyl,  $C_{2-6}$ alkynyl,  $C_{3-6}$ cycloalkyl, phenyl, naphthyl or a heterocyclic group, wherein in  $R^5$  any  $C_{1-6}$ alkyl,  $C_{2-6}$ alkenyl,  $C_{2-6}$ alkynyl,  $C_{3-6}$ cycloalkyl, phenyl, naphthyl or heterocyclic groups (on a ring carbon) may be optionally substituted by one or more  $R^6$  and if said heterocyclic group contains an NH moiety that nitrogen may be optionally substituted by a group selected from  $R^8$ ;

E is  $-O-$ ,  $N(R^9)-$ ,  $C(O)-$ ,  $N(R^9)C(O)-$ ,  $C(O)N(R^9)-$ ,  $S(O)_a-$  wherein  $a$  is 0-2,  $OC(O)-$ ,  $C(O)O-$ ,  $N(R^9)C(O)O-$ ,  $OC(O)N(R^9)-$ ,  $C(S)N(R^9)-$ ,  $N(R^9)C(S)-$ ,  $SO_2N(R^9)-$ ,  $N(R^9)SO_2-$ ,  $N(R^9)C(O)N(R^9)-$ ,  $N(R^9)C(S)N(R^9)-$ ,  $SO_2NHC(O)-$ ,  $SO_2N(R^9)C(O)-$ ,  $C(O)NHSO_2-$  or E is a direct bond;

**R<sup>6</sup>** is trifluoromethyl, C<sub>1-6</sub>alkyl, halo, hydroxy, trifluoromethoxy, cyano, C<sub>1-6</sub>alkoxy, formyl, C<sub>1-6</sub>alkanoyl, C<sub>1-6</sub>alkanoyloxy, amino, *N*-(C<sub>1-6</sub>alkyl)amino, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>amino, C<sub>1-6</sub>alkanoylamino, C<sub>1-6</sub>alkanoyl(*N*-C<sub>1-6</sub>alkyl)amino, nitro, carboxy, carbamoyl, C<sub>1-6</sub>alkoxycarbonyl, thiol, C<sub>1-6</sub>alkylsulphanyl, C<sub>1-6</sub>alkylsulphinyl, C<sub>1-6</sub>alkylsulphonyl, C<sub>1-6</sub>alkylsulphonylamino, sulphamoyl, *N*-(C<sub>1-6</sub>alkyl)aminosulphonyl, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>aminosulphonyl, *N*-(C<sub>1-6</sub>alkyl)carbamoyl, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>carbamoyl, ureido, *N'*-(C<sub>1-6</sub>alkyl)ureido or *N'*-(C<sub>1-6</sub>alkyl)<sub>2</sub>ureido, C<sub>2-6</sub>alkenyl, C<sub>2-6</sub>alkynyl, C<sub>3-6</sub>cycloalkyl, naphthyl; or phenyl or a heterocyclic group wherein in R<sup>6</sup> any C<sub>1-6</sub>alkyl, C<sub>2-6</sub>alkenyl, C<sub>2-6</sub>alkynyl, C<sub>3-6</sub>cycloalkyl ~~C<sub>3-6</sub>cycloalkyl~~, naphthyl; or phenyl or heterocyclic groups (on a ring carbon) may be optionally substituted by one or more R<sup>7</sup> ~~and if said heterocyclic group contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from R<sup>8</sup>~~;

**R<sup>7</sup>** is trifluoromethyl, cyano, C<sub>1-6</sub>alkyl, halo, hydroxy, trifluoromethoxy, C<sub>1-6</sub>alkoxy, formyl, C<sub>1-6</sub>alkanoyl, C<sub>1-6</sub>alkanoyloxy, amino, *N*-(C<sub>1-6</sub>alkyl)amino, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>amino, C<sub>1-6</sub>alkanoylamino, C<sub>1-6</sub>alkanoyl(*N*-C<sub>1-6</sub>alkyl)amino, nitro, carboxy, carbamoyl, C<sub>1-6</sub>alkoxycarbonyl, thiol, C<sub>1-6</sub>alkylsulphanyl, C<sub>1-6</sub>alkylsulphinyl, C<sub>1-6</sub>alkylsulphonyl, C<sub>1-6</sub>alkylsulphonylamino, sulphamoyl, *N*-(C<sub>1-6</sub>alkyl)aminosulphonyl, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>aminosulphonyl, *N*-(C<sub>1-6</sub>alkyl)carbamoyl, *N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>carbamoyl, C<sub>2-6</sub>alkenyl, C<sub>2-6</sub>alkynyl; or C<sub>3-6</sub>cycloalkyl or a heterocyclic group (optionally substituted by one or more R<sup>14</sup>), and wherein in R<sup>7</sup> any C<sub>1-6</sub>alkyl, C<sub>2-6</sub>alkenyl, C<sub>2-6</sub>alkynyl or C<sub>3-6</sub>cycloalkyl ~~C<sub>3-6</sub>cycloalkyl~~ groups may be optionally substituted by one or more groups selected from R<sup>12</sup>;

**R<sup>8</sup>** is ~~C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkanoyl, C<sub>1-6</sub>alkylsulphonyl, C<sub>1-6</sub>alkoxycarbonyl, carbamoyl, *N*-(C<sub>1-6</sub>alkyl)carbamoyl, *N,N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>carbamoyl, benzoyl, (heterocyclic group)carbonyl, phenylsulphonyl, (heterocyclic group)sulphonyl, phenyl or a carbon linked heterocyclic group, and wherein in R<sup>8</sup> any C<sub>1-6</sub>alkyl, phenyl or heterocyclic group (on a ring carbon) may be optionally substituted by one or more R<sup>6</sup>, and if a heterocyclic group contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from R<sup>14</sup>~~,

wherein for  $R^4$ ,  $R^5$ ,  $R^6$ ,  $R^7$  and  $R^8$ , a heterocyclic group is ~~selected from morpholino, piperidyl, pyridyl, pyranyl, pyrrolyl, isothiazolyl, indolyl, quinolyl, thienyl, 1,3-benzodioxolyl, thiadiazolyl, piperazinyl, thiazolidinyl, pyrrolidinyl, thiomorpholino, pyrrolinyl, homopiperazinyl, tetrahydropyranyl, imidazolyl, pyrimidyl, pyrazinyl, pyridazinyl, isoxazolyl, 4-pyridone, 1-isoquinolone, 2-pyrrolidone, 4-thiazolidone, pyridine *N*-oxide, quinoline *N*-oxide and combinations thereof;~~

$R^9$  is hydrogen or  $C_{1-6}$ alkyl optionally substituted by one or more  $R^{10}$  with the proviso that  $R^{10}$  is not a substituent on the carbon attached to a nitrogen atom;

$R^{10}$  is halo, hydroxy, amino, cyano, nitro, trifluoromethyl, trifluoromethoxy,  $C_{1-6}$ alkyl,  $C_{2-6}$ alkenyl,  $C_{2-6}$ alkynyl,  $N$ -( $C_{1-6}$ alkyl)amino,  $N$ -( $C_{1-6}$ alkyl)<sub>2</sub>amino,  $C_{1-6}$ alkanoylamino,  $C_{1-6}$ alkanoyl( $N$ - $C_{1-6}$ alkyl)amino,  $C_{1-6}$ alkylsulphonylamino,  $C_{1-6}$ alkylsulphonyl( $N$ - $C_{1-6}$ alkyl)amino, thiol,  $C_{1-6}$ alkylsulphanyl,  $C_{1-6}$ alkylsulphinyl,  $C_{1-6}$ alkylsulphonyl, sulphamoyl,  $N$ -( $C_{1-6}$ alkyl)aminosulphonyl,  $N$ -( $C_{1-6}$ alkyl)<sub>2</sub>aminosulphonyl, carboxy, carbamoyl,  $N$ -( $C_{1-6}$ alkyl)carbamoyl,  $N$ -( $C_{1-6}$ alkyl)<sub>2</sub>carbamoyl,  $C_{1-6}$ alkoxycarbonyl,  $C_{1-6}$ alkanoyl or formyl;

$R^{11}$  is  ~~$C_{1-6}$ alkyl,  $C_{1-6}$ alkanoyl,  $C_{1-6}$ alkylsulphonyl,  $C_{1-6}$ alkoxycarbonyl, carbamoyl,  $N$ -( $C_{1-6}$ alkyl)carbamoyl,  $N,N$ -( $C_{1-6}$ alkyl)<sub>2</sub>carbamoyl,  $C_{1-6}$ alkoxy $C_{1-6}$ alkanoyl, phenyl $C_{1-6}$ alkyl, benzoyl, phenyl $C_{1-6}$ alkanoyl, phenyl $C_{1-6}$ alkoxycarbonyl or phenylsulphonyl and wherein in  $R^{11}$  any  $C_{1-6}$ alkyl group can be optionally substituted by one or more  $R^{13}$ ;~~

$R^{12}$  is halo, hydroxy,  $N$ -methylpiperazinyl,  $N$ -acetylpiperazinyl, morpholino, piperidino, cyano, amino,  $N,N$ -dimethylamino, acetamido, carbamoyl, carboxy, methanesulphonyl or sulphamoyl;

$R^{13}$  is ~~halo, hydroxy, cyano, amino,  $N,N$ -dimethylamino, acetamido, carbamoyl, carboxy, methanesulphonyl or sulphamoyl;~~

$n$  is 0-5; wherein the values of  $R^3$  may be the same or different;

or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof.

Claim 2 (original): A compound of formula (I) according to claim 1 wherein one of  $R^1$  and  $R^2$  is methyl and the other is trifluoromethyl;

or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof.

Claim 3 (canceled).

Claim 4 (currently amended): A compound of formula (I) according to claim 1 wherein R<sup>3</sup> is a substituent on carbon and is selected from amino, and methyl, ~~4-mesylphenylsulphonyl, 4-methylthiophenylthio, 4-fluorobenzoyl and 4-cyanobenzoylamine;~~  
or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof.

Claim 5 (previously presented): A compound of formula (I) according to claim 1 wherein R<sup>4</sup> is C<sub>1-4</sub>alkyl, phenyl {optionally substituted with one or more *t*-butyl, isopropyl, nitro, halo, *N,N*-dimethylcarbamoyl, *N,N*-dimethylamino, 2-hydroxyethylamino, cyano, acetyl, methoxy or carboxy} or thienyl;  
or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof.

Claim 6 (previously presented): A compound of formula (I) according to claim 1 wherein D is -SO<sub>2</sub>-;  
or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof.

Claim 7 (previously presented): A compound of formula (I) according to claim 1 wherein n is 0 - 3;  
or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof.

Claim 8 (previously presented): A compound of formula (I) according to claim 1, selected from:

(R)-[(2S,5R)-2-methyl-5-methyl-4-(4-carboxyphenylsulphonyl)-1-(3,3,3-trifluoro-2-hydroxy-2-methylpropionyl)piperazine];

(R)-[(2S,5R)-2-methyl-5-methyl-4-(4-dimethylcarbamoylphenylsulphonyl)-1-(3,3,3-trifluoro-2-hydroxy-2-methylpropionyl)piperazine];

(R)-[(2S,5R)-2-methyl-5-methyl-4-(4-fluorophenylsulphonyl)-1-(3,3,3-trifluoro-2-hydroxy-2-methylpropionyl)piperazine];

(R)-{(2S,5R)-2-methyl-5-methyl-4-[4-(2-hydroxyethylamino)phenylsulphonyl]-1-(3,3,3-trifluoro-2-hydroxy-2-methylpropionyl)piperazine};

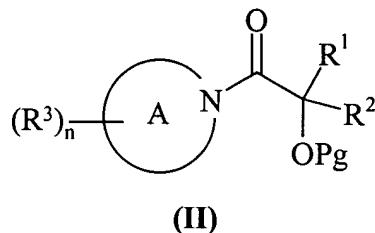
(R)-[(2S,5R)-2-methyl-5-methyl-4-(4-cyanophenylsulphonyl)-1-(3,3,3-trifluoro-2-hydroxy-2-methylpropionyl)piperazine]; and

(R)-[(2S,5R)-2-methyl-5-methyl-4-(4-methoxyphenylsulphonyl)-1-(3,3,3-trifluoro-2-hydroxy-2-methylpropionyl)piperazine];

or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof.

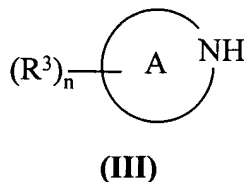
Claim 9 (previously presented): A process for preparing a compound of formula (I) as described in claim 1, or a pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, which process (in which variable groups are as defined in claim 1 for formula (I) unless otherwise stated) comprises of:

(a) deprotecting a protected compound of formula (II):

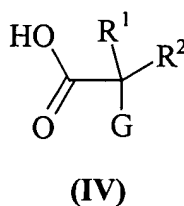


where Pg is an alcohol protecting group;

(b) coupling an amine of formula (III):



with an acid of formula (IV):



wherein G is a hydroxyl group;

(c) coupling an amine of formula (III) with an activated acid derivative of formula (IV)

wherein G is a hydroxyl group which may be protected as an ester or ether;

and thereafter if necessary:

i) converting a compound of the formula (I) into another compound of the formula (I);

ii) removing any protecting groups; or

iii) forming a pharmaceutically acceptable salt or *in vivo* hydrolysable ester.

Claim 10 (previously presented): A pharmaceutical composition which comprises a compound of formula (I) according to any one of claims 1-2 and 4-8, or a pharmaceutically acceptable salt or *in vivo* hydrolysable ester thereof in association with a pharmaceutically-acceptable diluent or carrier.

Claims 11-15 (canceled).

Claim 16 (previously presented): A method for the treatment of diabetes mellitus, said method comprising administering to a warm-blooded animal in need thereof a diabetes mellitus effective amount of a compound of the formula (I) or pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, as claimed in any one of claims 1-2 and 4-8.

Claim 17 (previously presented): A method for the treatment of peripheral vascular disease, said method comprising administering to a warm-blooded animal in need thereof a peripheral vascular disease effective amount of a compound of the formula (I) or pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, as claimed in any one of claims 1-2 and 4-8.

Claim 18 (currently amended): A method for the treatment of myocardial or cerebral ischaemia, said method comprising administering to a warm-blooded animal in need thereof a myocardial or cerebral ~~an~~-ischaemia effective amount of a compound of the formula (I) or

pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, as claimed in any one of claims 1-2 and 4-8.

Claim 19 (previously presented): A method for the treatment of hyperlipidaemia, said method comprising administering to a warm-blooded animal in need thereof a hyperlipidaemia effective amount of a compound of the formula (I) or pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, as claimed in any one of claims 1-2 and 4-8.

Claim 20 (currently amended): A method for the treatment of Alzheimer's ~~Alzheimers~~-disease, said method comprising administering to a warm-blooded animal in need thereof an Alzheimer's ~~Alzheimers~~-disease effective amount of a compound of the formula (I) or pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, as claimed in any one of claims 1-2 and 4-8.

Claim 21 (previously presented): A method for the treatment of atherosclerosis, said method comprising administering to a warm-blooded animal in need thereof an atherosclerosis effective amount of a compound of the formula (I) or pharmaceutically acceptable salt or an *in vivo* hydrolysable ester thereof, as claimed in any one of claims 1-2 and 4-8.